

# STIC Search Report Biotech-Chem Library

## STIC Database Tracking Number: 192170

TO: Marcela Cordero Garcia

Location: Rem/3a30/3c18

Art Unit: 1654

Tuesday, June 06, 2006

Case Serial Number: 10/796158

From: Toby Port

**Location: Biotech-Chem Library** 

**REM-1A59** 

Phone: (571)272-2523

toby.port@uspto.gov

### Search Notes

Dear Examiner Cordero Garcia.

See attached results.

If you have any questions about this search feel free to contact me at any time.

Thank you for using STIC search services!

Toby Port Technical Information Specialist STIC Biotech/Chem Library (571)272-2523





10/796,158

=> file reg; d que l1
FFEE: REGISTRY' ENTERED AT 16:04:18 ON 06 JUN 2006
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STRUCTURE FILE UPDATES: 5 JUN 2006 HIGHEST RN 886840-90-0 DICTIONARY FILE UPDATES: 5 JUN 2006 HIGHEST RN 886840-90-0

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TSCA INFORMATION NOW CURRENT THROUGH January 6, 2006

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Structure search iteration limits have been increased. See HELP SLIMITS for details.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

http://www.cas.org/ONLINE/UG/regprops.html

(L1 )

6 SEA FILE=REGISTRY ABB=ON PLU=ON YCYYCFWKTCT CYYYCFWKTCT YYCYC

=> /d ll rn cn sql kwic nte 1-6

L1 ANSWER 1 OF 6 REGISTRY COPYRIGHT 2006 ACS on STN

RN 760191-73-9 REGISTRY

CN L-Threonine, L-tyrosyl-L-tyrosyl-L-cysteinyl-L-tyrosyl-L-cysteinyl-Lphenylalanyl-L-tryptophyl-L-lysyl-L-threonyl-L-cysteinyl- (9CI) (CA INDEX
NAME)

OTHER NAMES:

CN 7: PN: WO2004081031 SEQID: 7 unclaimed sequence

SQL 11

SEQ 1 YYCYCFWKTC T

HITS AT: 1-11

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**RELATED SEQUENCES AVAILABLE WITH SEQLINK**
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ANSWER 2 OF 6 REGISTRY COPYRIGHT 2006 ACS on STN
L1
    760191-72-8 REGISTRY
RN
    L-Threonine, L-cysteinyl-L-tyrosyl-L-tyrosyl-L-cysteinyl-L-
CN
    phenylalanyl-L-tryptophyl-L-lysyl-L-threonyl-L-cysteinyl- (9CI) (CA INDEX
    NAME)
OTHER NAMES:
    6: PN: WO2004081031 SEQID: 6 unclaimed sequence
SOL 11
       1 CYYYCFWKTC T
SEO
         HITS AT:
         1-11
**RELATED SEOUENCES AVAILABLE WITH SEOLINK**
    ANSWER 3 OF 6 REGISTRY COPYRIGHT 2006 ACS on STN
L1
    760191-71-7 REGISTRY
RN
    L-Threonine, L-tyrosyl-L-cysteinyl-L-tyrosyl-L-tyrosyl-L-cysteinyl-L-
CN
    phenylalanyl-L-tryptophyl-L-lysyl-L-threonyl-L-cysteinyl- (9CI) (CA INDEX
OTHER NAMES:
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CN
SQL 11
SEO
       1 YCYYCFWKTC T
         HITS AT:
         1-11
**RELATED SEOUENCES AVAILABLE WITH SEOLINK**
L1
    ANSWER 4 OF 6 REGISTRY COPYRIGHT 2006 ACS on STN
    757951-73-8 REGISTRY
RN
    L-Threonine, D-tyrosyl-D-tyrosyl-D-cysteinyl-D-tyrosyl-L-cysteinyl-D-
CN
    phenylalanyl-L-tryptophyl-L-lysyl-L-threonyl-L-cysteinyl-, cyclic
    (5→10)-disulfide (9CI) (CA INDEX NAME)
SOL
    11
SEO
       1 YYCYCFWKTC T
         HITS AT:
         1-11
**RELATED SEOUENCES AVAILABLE WITH SEOLINK**
______
            ----- location ----- description
type
_____
bridge Cys-5 - Cys-10 disulfide bridge
______
L1
    ANSWER 5 OF 6 REGISTRY COPYRIGHT 2006 ACS on STN
RN
    757951-72-7 REGISTRY
    L-Threonine, D-cysteinyl-D-tyrosyl-D-tyrosyl-D-tyrosyl-L-cysteinyl-D-
CN
    phenylalanyl-L-tryptophyl-L-lysyl-L-threonyl-L-cysteinyl-, cyclic
    (5→10) -disulfide (9CI) (CA INDEX NAME)
SQL 11
SEQ
       1 CYYYCFWKTC T
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HITS AT: 1-11

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

NTE

type ----- location ----- description

bridge Cys-5 - Cys-10 disulfide bridge

L1 ANSWER 6 OF 6 REGISTRY COPYRIGHT 2006 ACS on STN

RN 757951-71-6 REGISTRY

CN L-Threonine, D-tyrosyl-D-cysteinyl-D-tyrosyl-D-tyrosyl-L-cysteinyl-Dphenylalanyl-L-tryptophyl-L-lysyl-L-threonyl-L-cysteinyl-, cyclic
(5→10)-disulfide (9CI) (CA INDEX NAME)

SOL 11

SEQ 1 YCYYCFWKTC T

HITS AT: 1-11

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

NTE

type ----- location ----- description

bridge Cys-5 - Cys-10 disulfide bridge

=> file caplus; d que 12

FILE 'CAPLUS' ENTERED AT 16:07:50 ON 06 JUN 2006

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L1 6 SEA FILE=REGISTRY ABB=ON PLU=ON YCYYCFWKTCT CYYYCFWKTCT YYCYC

L2 1 SEA FILE=CAPLUS ABB=ON PLU=ON L1

#### => d ibib ed ab hitrn

L2 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:780722 CAPLUS

DOCUMENT NUMBER: 141:271609

TITLE: Thiol-mediated drug attachment to targeting peptides

INVENTOR(S): Braslawsky, Gary R.; Chinn, Paul

\_\_\_\_

PATENT ASSIGNEE(S): Biogen Idec Inc., USA SOURCE: PCT Int. Appl., 43 pp. CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PA.	CENT 1	NO.			KIND DATE					APPL	ICAT:	I NOI	. O <i>l</i>	DATE				
						-						<del>-</del>						
WO	2004	0810	31		A2		2004	923	1	WO 2	004-1	JS714	20040310					
WO	2004	0810	31		<b>A</b> 3		2005	0310										
	W:	ΑE,	AG,	AL,	AM,	AT,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BW,	BY,	ΒZ,	CA,	CH,	
		CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,	
		GE,	GH,	GM,	HR,	ΗU,	ID,	ΙL,	IN,	IS,	JP,	KΕ,	KG,	ΚP,	KR,	KΖ,	LC,	
		LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	ΜZ,	NA,	NI,	
		NO,	ΝZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,	
		ТJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UΖ,	VC,	VN,	YU,	ZA,	ZM,	ZW	
	RW:	BW,	GH,	GM,	KE,	LS,	MW,	ΜZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	ΑZ,	
		BY,	KG,	KZ,	MD,	RU,	ТJ,	TM,	AT,	ΒE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	
		ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC,	NL,	PL,	PT,	RO,	SE,	SI,	
		SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	NΕ,	SN,	
		TD,	TG															
ΑŬ	20042	2201	04		A1		2004	0923	1	AU 2	004-3	2201		20	0040	310		
CA	25184	406			AA		2004	923	(	CA 2	004-2	25184	406		20	00403	310	
US	2005	1180	99		A1		2005	0602	Ī	US 2	004-'	7961	58		20	00403	310	
EP	16108	805			A2		2006	0104	1	EP 2	004-	7191	92		20	0403	310	
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,	
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PRIORITY APPLN. INFO.:				. :					US 2003-452928P					P 20030310				
									1	WO 2	004-1	JS714	A 20040310					

ED Entered STN: 24 Sep 2004

AB The invention discloses compns. and methods for thiol-specific attachment of therapeutic and diagnostic agents to somatostatin and other targeting peptides. Compns. of the invention include somatostatin analogs AB (A = cysteine or cysteine-containing peptide suitable for binding to drug or chelator via thiol linkage; B = somatostatin peptide or fragment that binds to somatostatin receptor).

TT 757951-71-6D, C-terminus amide or alc. 757951-72-7D,
 C-terminus amide or alc. 757951-73-8D, C-terminus amide or alc.
 RL: DGN (Diagnostic use); PAC (Pharmacological activity); PRP
 (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (thiol-mediated drug attachment to targeting peptides)

IT 760191-71-7 760191-72-8 760191-73-9

RL: PRP (Properties)

(unclaimed sequence; thiol-mediated drug attachment to targeting peptides)

=> => file caplus; d que 15 FILE 'CAPLUS' ENTERED AT 16:26:43 ON 06 JUN 2006 10/796,158

Cordero Garcia

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L3	41 8	SEA	FILE=CAPLUS	ABB=ON	PLU=ON	BRASLAWSKY G?/AU
L4	21 \$	SEA	FILE=CAPLUS	ABB=ON	PLU=ON	CHINN P?/AU
L5	5 5	SEA	FILE=CAPLUS	ABB=ON	PLU=ON	(L3 OR L4) AND PEPTIDES/CW

=> file medline; d que 18 FILE MEDLINE' ENTERED AT 16:27:01 ON 06 JUN 2006

FILE LAST UPDATED: 3 JUN 2006 (20060603/UP). FILE COVERS 1950 TO DATE.

On December 11, 2005, the 2006 MeSH terms were loaded.

The MEDLINE reload for 2006 is now (26 Feb.) available. For details on the 2006 reload, enter HELP RLOAD at an arrow prompt (=>). See also:

http://www.nlm.nih.gov/mesh/

http://www.nlm.nih.gov/pubs/techbull/nd04/nd04 mesh.html

http://www.nlm.nih.gov/pubs/techbull/nd05/nd05 med\_data\_changes.html

http://www.nlm.nih.gov/pubs/techbull/nd05/nd05 2006 MeSH.html

OLDMEDLINE is covered back to 1950.

L8

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2006 vocabulary.

This file contains CAS Registry Numbers for easy and accurate substance identification.

41 SEA FILE=CAPLUS ABB=ON PLU=ON BRASLAWSKY G?/AU L321 SEA FILE=CAPLUS ABB=ON PLU=ON CHINN P?/AU L4

SEA FILE=MEDLINE ABB=ON PLU=ON (L3 OR L4) AND PEPTIDE?

=> file\_embase; d que l12 FILE "EMBASE' ENTERED AT 16:27:08 ON 06 JUN 2006 Copyright (c) 2006 Elsevier B.V. All rights reserved.

FILE COVERS 1974 TO 6 Jun 2006 (20060606/ED)

EMBASE has been reloaded. Enter HELP RLOAD for details.

EMBASE is now updated daily. SDI frequency remains weekly (default) and biweekly.

This file contains CAS Registry Numbers for easy and accurate substance identification.

41 SEA FILE=CAPLUS ABB=ON PLU=ON BRASLAWSKY G?/AU L3 L4\_\_\_\_\_

21 SEA FILE=CAPLUS ABB=ON PLU=ON CHINN P?/AU

/2 SEA FILE=EMBASE ABB=ON PLU=ON (L3 OR L4) AND PEPTIDE? 〔L12... ... ≥

=> file biosis; d que l13 FILE BIOSIS ENTERED AT 16:27:15 ON 06 JUN 2006 Copyright (c) 2006 The Thomson Corporation

FILE COVERS 1969 TO DATE. CAS REGISTRY NUMBERS AND CHEMICAL NAMES (CNs) PRESENT FROM JANUARY 1969 TO DATE.

RECORDS LAST ADDED: 31 May 2006 (20060531/ED)

L3 L4 1613 41 SEA FILE=CAPLUS ABB=ON PLU=ON BRASLAWSKY G?/AU
21 SEA FILE=CAPLUS ABB=ON PLU=ON CHINN P?/AU
2 SEA FILE=BIOSIS ABB=ON PLU=ON (L3 OR L4) AND PEPTIDE?

file wpix; d que 114

FINE WPIX' ENTERED AT 16:27:21 ON 06 JUN 2006

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FILE LAST UPDATED: 2 JUN 2006 <20060602/UP>
MOST RECENT DERWENT UPDATE: 200635 <200635/DW>
DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE

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 PLEASE VISIT:
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L3 41 SEA FILE=CAPLUS ABB=ON PLU=ON BRASLAWSKY G?/AU
L4 21 SEA FILE=CAPLUS ABB=ON PLU=ON CHINN P?/AU
L14 7 SEA FILE=WPIX ABB=ON PLU=ON (L3 OR L4) AND PEPTIDE?

=> dup rem 18 15 112 113 114 FILE 'MEDLINE' ENTERED AT 16:27:36 ON 06 JUN 2006

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PROCESSING COMPLETED FOR L8
PROCESSING COMPLETED FOR L5
PROCESSING COMPLETED FOR L12
PROCESSING COMPLETED FOR L13
PROCESSING COMPLETED FOR L14
L15
12 DUP REM L8 L5 L12 L13 L14 (7 D

12 DUP REM L8 L5 L12 L13 L14 (7 DUPLICATES REMOVED)
ANSWERS '1-3' FROM FILE MEDLINE
ANSWERS '4-8' FROM FILE CAPLUS
ANSWERS '9-12' FROM FILE WPIX

=> d ibib ed ab 115 1-8; d ibib ab abex 115 9-12

L15 ANSWER 1 OF 12 MEDLINE on STN DUPLICATE 3

ACCESSION NUMBER: 93294279 MEDLINE DOCUMENT NUMBER: PubMed ID: 8515057

TITLE: Identification of functional domains in murine

granulocyte-macrophage colony-stimulating factor using

monoclonal antibodies to synthetic **peptides**. Greenfield R S; **Braslawsky G R**; Kadow K F; Spitalny G L; Chace D; Bull C O; Bursuker I

CORPORATE SOURCE: Bristol-Myers Squibb Co., Pharmaceutical Research

Institute, Wallingford, CT 06492.

SOURCE: Journal of immunology (Baltimore, Md.: 1950), (1993 Jun

15) Vol. 150, No. 12, pp. 5241-51.

Journal code: 2985117R. ISSN: 0022-1767.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals

ENTRY MONTH: 199307

**AUTHOR:** 

ENTRY DATE: Entered STN: 6 Aug 1993

Last Updated on STN: 6 Aug 1993 Entered Medline: 20 Jul 1993

ED Entered STN: 6 Aug 1993

Last Updated on STN: 6 Aug 1993 Entered Medline: 20 Jul 1993

Granulocyte-macrophage (GM)-CSF is an important hematopoietic cytokine AB that regulates proliferation and differentiation of macrophages, neutrophils, and eosinophils. In this study, we generated mAb to five synthetic peptides that correspond to regions along the murine GM-CSF molecule. The ability of anti-peptide mAb to bind to and inhibit biologic activity of murine (m) GM-CSF was determined. mAb with the highest neutralization titers were derived from mice immunized with peptide II, which correspond to amino acids 27 to 38 of mGM-CSF. Immunochemical studies showed that peptide II specifically blocked binding of anti-peptide II mAb to GM-CSF. mAb to two other peptides in the N-terminal half corresponding to residues 7 to 17 and 47 to 58, respectively, of mGM-CSF also inhibited GM-CSF-dependent proliferation and differentiation of murine bone marrow precursors for macrophages and granulocytes. Anti-peptide mAb also inhibited growth of a murine hematopoietic cell line FDCP1 and a murine T cell line HT-2, which was shown to be dependent on GM-CSF for growth in vitro. Biologic activity of both natural and recombinant mGM-CSF was neutralized by anti-peptide mAb. These findings indicate that epitopes in the N-terminal region of mGM-CSF are important for biologic activity, and the epitope defined by peptide II (residues 27 to 38) lies within a particularly important functional domain of the mGM-CSF molecule.

L15 ANSWER 2 OF 12 MEDLINE on STN DUPLICATE 4

ACCESSION NUMBER: 94065479 MEDLINE DOCUMENT NUMBER: PubMed ID: 8245704

TITLE: Granulocyte-macrophage colony-stimulating factor plays a

role in the functional activity of mast cells.

AUTHOR: Meade R; Neddermann K M; Greenfield R S; Braslawsky

G; Bursuker I

CORPORATE SOURCE: Bristol-Myers Squibb Pharmaceutical Research Institute,

Wallingford, CT 06492-7660.

SOURCE: Journal of leukocyte biology, (1993 Dec) Vol. 54, No. 6,

pp. 523-7.

Journal code: 8405628. ISSN: 0741-5400.

10/796,158 Cordero Garcia

United States PUB. COUNTRY:

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

English LANGUAGE:

Priority Journals FILE SEGMENT:

ENTRY MONTH: 199401

ENTRY DATE: Entered STN: 1 Feb 1994

Last Updated on STN: 1 Feb 1994

Entered Medline: 6 Jan 1994

ED Entered STN: 1 Feb 1994

Last Updated on STN: 1 Feb 1994

Entered Medline: 6 Jan 1994

AΒ A peptide homologous to a region of murine granulocytemacrophage colony-stimulating factor (mGM-CSF), P27-38, which was shown to be a GM-CSF antagonist, inhibited the function of serotonin release from murine mast cells. Peptide P27-38 inhibited immunoglobulin E (IgE) -mediated serotonin release in a dose-dependent manner when induced by either specific antiqen or anti-IqE antibody. In contrast, non-receptor-mediated release of serotonin by agents such as compound 48/80 or the calcium ionophore A23187 were not affected by the GM-CSF antagonist. Similar effects were observed with GM-CSF-neutralizing antibodies. The inhibitory effect of P27-38 and the neutralizing antibodies on serotonin release could be reversed by the addition of exogenous GM-CSF to the stimulated mast cells, indicating that the inhibitory activity was probably due to an effect on endogenously produced GM-CSF. These findings suggest that GM-CSF produced by stimulated mast cells is involved in the regulation of their activity in an autocrine manner.

L15 ANSWER 3 OF 12 MEDLINE on STN ACCESSION NUMBER: 85238871 MEDLINE DOCUMENT NUMBER: PubMed ID: 2409352

TITLE: Effect of a cyclic hexapeptide analog (L363,586) of

somatostatin on the function of pancreas grafts in dogs.

AUTHOR: Liu T; Sutherland D E; Chinn P L; Najarian J S

The Journal of surgical research, (1985 Jul) Vol. 39, No. SOURCE:

1, pp. 39-45.

Journal code: 0376340. ISSN: 0022-4804.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

Priority Journals FILE SEGMENT:

ENTRY MONTH: 198508

Entered STN: 20 Mar 1990 ENTRY DATE:

> Last Updated on STN: 6 Feb 1995 Entered Medline: 20 Aug 1985

ED Entered STN: 20 Mar 1990

> Last Updated on STN: 6 Feb 1995 Entered Medline: 20 Aug 1985

Complications related to the exocrine secretions cause some pancreas AΒ grafts to fail in the early postoperative period. Somatostatin inhibits exocrine secretion, as well as insulin and glucagon release. L363,586 is a cyclic hexapeptide analog of somatostatin that is 50 to 100 times more potent than the native hormone in inhibiting islet hormone release. In a preliminary experiment in which permanent fistulas were created in two dogs, we demonstrated that L363,586 (0.3 micrograms/kg/60 min) results in a fourfold decrease in pancreatic exocrine secretion when measured for 210 min following a beef meal. In a separate experiment, five totally pancreatectomized dogs who received segmental pancreas autografts with pancreaticoductocystostomy 10 months previously had L363,586 (0.3 micrograms/kg/hr) administered by the Alzet osmotic pump subcutaneously

for 7 days. Mean (+/-SE) daily serum amylase activity (IU/dl) during the week before the implant was 78 +/- 3, during the week of infusion was 65 +/- 2 (P less than 0.001), and during the week afterward was 76 +/- 2. In a prospective experiment, 12 totally pancreatectomized dogs received segmental pancreas autografts with anastomosis of the graft vessels to the iliac vessels and of the pancreatic duct to the bladder. L363,586 was administered by osmotic pump for 7 days to seven dogs at a dose of 0.3 micrograms/kg/hr. Mean (+/-SE) daily serum amylase levels at 1, 2, and 3 weeks posttransplant were 223 +/- 17, 81 +/- 3, and 82 +/- 5 in the L363,586-treated dogs and 229 +/- 18, 108 +/- 5, and 90 +/- 5 in the five untreated dogs (P less than 0.001 at 2 weeks). (ABSTRACT TRUNCATED AT 250 WORDS)

L15 ANSWER 4 OF 12 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 1

ACCESSION NUMBER: 2004:780722 CAPLUS

DOCUMENT NUMBER: 141:271609

TITLE: Thiol-mediated drug attachment to targeting peptides

INVENTOR(S): Braslawsky, Gary R.; Chinn, Paul

PATENT ASSIGNEE(S): Biogen Idec Inc., USA SOURCE: PCT Int. Appl., 43 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND DATE	APPLICATION NO.	DATE
WO 2004081031 WO 2004081031	A2 20040923	WO 2004-US7143	20040310
W: AE, AG, AL,	AM, AT, AU, AZ,	BA, BB, BG, BR, BW, 1	BY, BZ, CA, CH,
CN. CO. CR.	CU. CZ. DE. DK.	DM, DZ, EC, EE, EG, 1	ES. FI. GB. GD.
, , ,		IN, IS, JP, KE, KG,	
, , ,			
		MD, MG, MK, MN, MW, I	
NO, NZ, OM,	PG, PH, PL, PT,	RO, RU, SC, SD, SE,	SG, SK, SL, SY,
TJ, TM, TN,	TR, TT, TZ, UA,	UG, US, UZ, VC, VN,	YU, ZA, ZM, ZW
		SD, SL, SZ, TZ, UG,	
		AT, BE, BG, CH, CY, C	
ES, FI, FR,	GB, GR, HU, IE,	IT, LU, MC, NL, PL,	PT, RO, SE, SI,
SK, TR, BF,	BJ, CF, CG, CI,	CM, GA, GN, GQ, GW, I	ML, MR, NE, SN,
TD, TG			
•	71 20040922	AU 2004-220104	20040310
CA 2518406			20040310
US 2005118099	A1 20050602	US 2004-796158	20040310
EP 1610805	A2 20060104	EP 2004-719192	20040310
		GB, GR, IT, LI, LU, I	
	• • • • • •		
	LV, FI, RU, MK,	CY, AL, TR, BG, CZ, I	
PRIORITY APPLN. INFO.:		US 2003-452928P	P 20030310
	•	WO 2004-US7143	A 20040310

ED Entered STN: 24 Sep 2004

The invention discloses compns. and methods for thiol-specific attachment of therapeutic and diagnostic agents to somatostatin and other targeting peptides. Compns. of the invention include somatostatin analogs AB (A = cysteine or cysteine-containing peptide suitable for binding to drug or chelator via thiol linkage; B = somatostatin peptide or fragment that binds to somatostatin receptor).

L15 ANSWER 5 OF 12 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 2

ACCESSION NUMBER: 2000:628157 CAPLUS

DOCUMENT NUMBER: 133:219807

10/796,158 Cordero Garcia

TITLE: Kit for radiolabeling proteins with yttrium-90

INVENTOR(S): Chinn, Paul

PATENT ASSIGNEE(S): Idec Pharmaceuticals Corporation, USA

SOURCE: PCT Int. Appl., 48 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	PATENT NO.					אדאום האידי				ADDITORTON NO								האתים			
	PAT	ENT.	NO.			VTM:		DAIE													
	MO	2000	0520°							WO 2000-US5078								2000	0229		
		2000									•••	20		3330	, 0			2000	0227		
	WO										BC	1	BR	RY	CA	СН	CN	CR	, CU,		
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	CA	2362	•	•	•			2000										2000	0229		
	CA 2362908 NZ 513667				A 20010928				NZ 2000-513667								2000	0229			
	EP 1156835			A2		2001	1128		ΕP	20	00-9	9193	45			2000	0229				
		R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GF	۲,	IT,	LI,	LU,	NL,	SE	, MC	, PT,		
			ΙE,	SI,	LT,	LV,	FI,	RO													
	BR	2000	0086	35		Α		2001	1226		BR	20	00-8	8635				2000	0229		
		2001							1015									2000	0229		
	JP	2002	5381	64		T2		2002	1112	JP 2000-602256							20000229				
	RU	2221	807			C2		2004	0120		RU	20	01-3	1263	96			2000	0229		
	ΑU	7803	11			B2		2005	0317									2000	0229		
	US	6994	840			B1		2006	0207	•	US	20	00-6	6281	86			2000	0728		
	za	2001	0069					2002	1122									2001	0822		
		1058				Α		2002			BG	20	01-3	1058	53			2001			
		2001						2001	1101									2001			
		2001																2001			
		2006				A1		2006	0330									2005			
PRIO	RITY	APP	LN.	INFO	.:													1999			
															78			2000			
				_						·	US	20	00-6	5281	86		A1	2000	0728		

ED Entered STN: 10 Sep 2000

AB Methods and kits for radiolabeling proteins and peptides with radiolytic isotopes, particularly yttrium-90, are disclosed, whereby sufficient purity, specific activity and binding affinity are achieved such that the radiolabeled protein may be directly administered to a patient without further column purification Such kits and methods will be particularly useful in bringing radioimmunotherapy to the hospital and outpatient setting for the treatment of cancer.

L15 ANSWER 6 OF 12 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 5

ACCESSION NUMBER: 1991:499410 CAPLUS

DOCUMENT NUMBER: 115:99410

TITLE: Anthracycline derivative conjugates having a novel

linker, methods for their production, and their use as

cytotoxic agents for targeting therapy

INVENTOR(S): Greenfield, Robert S.; Braslawsky, Gary R.;

Olech, Lee J.; Kaneko, Takushi; Kiener, Peter A.

PATENT ASSIGNEE(S): Bristol-Myers Co., USA SOURCE: Eur. Pat. Appl., 70 pp.

CODEN: EPXXDW

DOCUMENT TYPE: LANGUAGE: Patent English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 398305	A2	19901122	EP 1990-109268	19900516
EP 398305	A3	19910320		
EP 398305	B1	19970319		
R: AT, BE, CH,	DE, DK	, ES, FR, G	BB, GR, IT, LI, LU, NL,	SE
US 5122368	Α	19920616	US 1989-353729	19890517
CA 2016584	AA	19901117	CA 1990-2016584	19900511
CA 2016584	С	19990629		
IL 94379	A1	19970218	IL 1990-94379	19900514
FI 102356	B1	19981130	FI 1990-2387	19900514
NO 9002197	Α	19901119	NO 1990-2197	19900516
NO 300691	B1	19970707		
AU 9055117	A1	19901122	AU 1990-55117	19900516
AU 631638	B2	19921203		
ZA 9003757	A	19920129	ZA 1990-3757	19900516
AT 150321	E	19970415	AT 1990-109268	19900516
ES 2099075	T3	19970516	ES 1990-109268	19900516
JP 03027321	A2	19910205	JP 1990-125629	19900517
JP 3062696	B2	20000712		
KR 136899	B1	19980425	KR 1990-7085	19900517
PRIORITY APPLN. INFO.:			US 1989-353729	A 19890517
			US 1988-155181	B2 19880211
			US 1988-270509	B2 19881116

OTHER SOURCE(S): MARPAT 115:99410

ED Entered STN: 06 Sep 1991

The title conjugates are provided, as are methods for their production, AB pharmaceutical compns., and methods for delivering cytotoxic anthracyclines to a selected population of cells desired to be eliminated. The anthracycline conjugates comprise ≥1 anthracycline mol. linked to a ligand that is reactive with a cell population to be eliminated, the anthracycline having a keto group at the C-13 position, and being attached to the ligand via a linker arm and being bound to that linker arm via an acid-sensitive acylhydrazone bond at the 13-keto position of the anthracycline. The conjugates of the invention are therefore useful in antibody- or ligand-mediated drug delivery systems for the preferential killing of a selected cell population in the treatment of diseases such as cancers and other tumors non-cytocidal viral or other pathogenic infections, and autoimmune disorders. Thus, a cysteine-containing bombesin analog was synthesized, purified, and reacted with adriamycin 13-[3-(2-pyridyldithio)propionyl]hydrazone-HCl (I) (preparation given). Binding activity of the peptide in the bombesin-adriamycin conjugate was not disturbed, the conjugate retaining the ability to bond to bombesin receptor-pos. cells. The conjugate was highly cytotoxic toward SVT2 transformed fibroblast cells and was more potent than free adriamycin. portion of the cytotoxic activity of the conjugate was blocked by excess bombesin. The conjugate was also specifically cytotoxic toward HCT116 colon carcinoma cells and Swiss 3T3 cells. Conjugates of adriamycin with various monoclonal antibodies, with EGF, and with transferrin are also described.

L15 ANSWER 7 OF 12 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1993:641371 CAPLUS

DOCUMENT NUMBER: 119:241371

TITLE: Thioether-linked drug-ligand conjugates

INVENTOR(S):
Willner, David; Trail, Pamela A.; King, Dalton H.;

Hofstead, Sandra J.; Greenfield, Robert S.;

Braslawsky, Gary R.

PATENT ASSIGNEE(S): Bristol-Myers Squibb Co., USA

SOURCE: Eur. Pat. Appl., 66 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE	
EP 554708 EP 554708	A1				
EP 554708	B1	20050504			
R: AT, BE, CH	, DE, DK	, ES, FR, C	GB, GR, IE, IT, LI, I	JU, MC, NL, PT, SE	
US 5622929	Α	19970422	US 1992-824951	19920123	
CA 2087286	AA	19930724	US 1992-824951 CA 1993-2087286	19930114	
CA 2087286	С	20040406			
AT 294592	E	20050515	AT 1993-100732	19930119	
ES 2240959	Т3	20051016	ES 1993-100732 AU 1993-31881	19930119	
AU 9331881	A1	19930729	AU 1993-31881	19930120	
AU 666903	B2	19960229			
ZA 9300444	Α	19930721	ZA 1993-444	19930121	
NO 9300189	Α		NO 1993-189		
JP 06025012 HU 68345	A2	19940201	JP 1993-40372	19930121	
HU 68345	A2	19950628	HU 1993-156	19930121	
RO 112618		19971128		19930121	
PL 172718	B1	19971128	PL 1993-317516	19930122	
PL 172715			PL 1993-317519		
PL 172828 PL 172837	B1	19971231	PL 1993-297514	19930122	
			PL 1993-317517		
PL 172827	B1		PL 1993-317518		
PL 172824		19971231	PL 1993-317715		
CN 1074684	Α	19930728	CN 1993-100709	19930123	
CN 1040540	В	19981104			
US 5606017	Α	19970225	US 1995-468162		
US 5708146	A A	19980113	US 1995-469840		
CN 1207946	Α	19990217	CN 1997-117785		
	Α	19980506	CN 1997-117908		
PRIORITY APPLN. INFO.:			US 1992-824951	A 19920123	
OTHER SOURCE(S):	MARPAT	119:24137	1		

OTHER SOURCE(S): MARPAT 119:241371

ED Entered STN: 11 Dec 1993

AB Drug-ligand conjugates [D=NNHCO(CH2)nAS((CH2)pC(=Y)NH)z]qX (D = drug; n = 1-10; p = 1-6; Y = 0, NH2+Cl-; z = 0, 1; q = 1-10; X = ligand; A = Michael addition adduct) are prepared for therapeutic use. Adriamycin hydrochloride was reacted with maleimidocaproyl hydrazide (preparation given) and then conjugated with thiolated monoclonal antibodies (MAbs), reduced MAbs, or modified bombesin [(Cys0, Lys3)bombesin]. The conjugates had antitumor activity in mice.

L15 ANSWER 8 OF 12 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1992:610646 CAPLUS

DOCUMENT NUMBER: 117:210646

TITLE: GM-CSF-inhibiting oligopeptides

INVENTOR(S):
Bursuker, Isia; Greenfield, Robert S.;

Braslawsky, Gary R.

PATENT ASSIGNEE(S): Bristol-Myers Squibb Co., USA

SOURCE: Eur. Pat. Appl., 22 pp.

10/796,158 Cordero Garcia

CODEN: EPXXDW

DOCUMENT TYPE: LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT:

1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE			
EP 499162	A2	19920819	EP 1992-102105	19920207			
EP 499162	A3	19930407					
R: AT, BE, CH,	DE, DK	, ES, FR, GB	, GR, IT, LI, LU, MC,	NL, PT, SE			
CA 2060699	AA	19920812	CA 1992-2060699	19920205			
JP 05239093	A2	19930917	JP 1992-65472	19920206			
PRIORITY APPLN. INFO.:			US 1991-653427	A 19910211			

ED Entered STN: 28 Nov 1992

Oligopeptides (sequences included) are provided which can inhibit the AR biol. activity of GM-CSF. The oligopeptides are useful for alleviating undesirable biol. effects mediated by GM-CSF. In particular, the oligopeptides are useful for inhibiting the growth of a GM-CSF-dependent neoplastic disease. Also provided are formulations and methods for using the oligopeptides. Murine GM-CSF-derived peptide Asp-Asp-Met-Pro-Val-Thr-Leu-Asn-Glu-Glu-Val-Glu inhibited, in a dose-dependent manner, the GM-CSF-induced proliferation of HT-2 cells.

L15 ANSWER 9 OF 12 WPIX COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER:

2005-058133 [06] WPIX

CROSS REFERENCE: DOC. NO. CPI:

2005-058132 [06] C2005-020196

TITLE:

New composition comprising polypeptide dimers comprising at least four binding sites and at least two polypeptide

chains linked via at least one interchain disulfide linkage, useful for treating e.g., cancer or autoimmune

diseases.

DERWENT CLASS:

B04 D16

INVENTOR(S):

CHINN, P; GLASER, S; REFF, M; WU, X; YANG, T

PATENT ASSIGNEE(S):

(BIOJ) BIOGEN IDEC MA INC

COUNTRY COUNT:

109

PATENT INFORMATION:

PATENT NO	KIND DATE	WEEK	LA PG
	<b></b>		

WO 2005000899 A2 20050106 (200506) \* EN 172

RW: AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IT KE LS LU MC MW MZ NA NL OA PL PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG

KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NA NI NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL SY TJ TM TN TR TT TZ UA UG

US UZ VC VN YU ZA ZM ZW

US 2005163782 A1 20050728 (200550)

EP 1641827 A2 20060405 (200624) EN

R: AL AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IT LI LT LU LV MC MK NL PL PT RO SE SI SK TR

#### APPLICATION DETAILS:

PATENT NO KIND APPLICATION DATE

10/796,158	Cordero Garcia

					- <b></b>	
WO	2005000899	A2		WO	2004-US20945	20040628
US	2005163782	<b>A1</b>	Provisional	US	2003-483877P	20030627
			Provisional	US	2003-508810P	20031003
			Provisional	US	2003-515351P	20031028
			Provisional	US	2003-516030P	20031030
				US	2004-880028	20040628
ΕP	1641827	<b>A2</b>		EP	2004-777279	20040628
				WO	2004-US20945	20040628

#### FILING DETAILS:

PATENT NO	KIND	PATENT NO
EP 1641827	A2 Based on	WO 2005000899
PRIORITY APPLN.	INFO: US 2003-516030P 2003-483877P 2003-508810P 2003-515351P 2004-880028	20031030; US 20030627; US 20031003; US 20031028; US 20040628

AB WO2005000899 A UPAB: 20060410

NOVELTY - A composition comprising polypeptide dimers comprising at least four binding sites and at least two polypeptide chains, where the polypeptide chains comprise at least one heavy chain portion and a synthetic connecting **peptide**, and where greater than about 50% of the dimers comprise polypeptide chains that are linked via at least one interchain disulfide linkage, is new.

DETAILED DESCRIPTION - A composition comprising polypeptide dimers comprising at least four binding sites and at least two polypeptide chains, where the polypeptide chains comprise at least one heavy chain portion and a synthetic connecting **peptide**, and where greater than about 50% of the dimers comprise polypeptide chains that are linked via at least one interchain disulfide linkage, or comprising minibody molecules comprising two polypeptide chains, where the polypeptide chains comprise a heavy chain portion and a synthetic connecting **peptide**, where the polypeptide chains lack all or part of a CH2 domain, and where greater than about 50% of the molecules are present in a form in which one of the polypeptide chains are linked via at least one interchain disulfide linkage, is new. INDEPENDENT CLAIMS are also included for:

- (1) a nucleic acid molecule comprising a nucleotide sequence encoding a polypeptide chain as defined above, or comprising a nucleotide sequence comprising 1857, 663, 1980, 1866, 645, 1173, or 1173 (SEQ ID NO: 17, 18, 23, 26, 27, 30 or 31, respectively) fully defined in the specification;
  - (2) a host cell comprising a vector; and
- (3) a binding molecule comprising the amino acid sequence comprising 621, 220, 661, 621, 214, 390 or 390 (SEQ ID NO: 20, 21, 25,28, 29, 32 or 33, respectively) fully defined in the specification.

ACTIVITY - Cytostatic; Immunosuppressive; Antiinflammatory; Gastrointestinal-Gen.; Dermatological; Antiulcer; Antirheumatic; Antiarthritic; Nephrotropic; Antithyroid; Thyromimetic; Muscular-Gen.; Neuroprotective; Antianemic; CNS-Gen.; Respiratory-Gen.; Vulnerary. No biological data given.

MECHANISM OF ACTION - None given.

USE - The composition is useful for treating a subject that would benefit from treatment with an antigen binding molecule, where the subject is suffering from cancer, lymphoma, an autoimmune disease or disorder, or an inflammatory disease or disorder (claimed). The composition is useful for treating autoimmune diseases such as Crohn's disease, inflammatory bowel disease, systemic lupus erythematosus, ulcerative colitis,

#### Cordero Garcia

rheumatoid arthritis, Goodpasture's syndrome, Grave's disease, Hashimoto's thyroiditis, pemphigus vulgaris, myasthenia gravis, scleroderma, autoimmune hemolytic anemia, pernicious anemia, Sjogren's syndrome, neurological disorders such as multiple sclerosis, and inflammatory diseases or disorders such as cystic fibrosis, sinusitis, gastroenteritis, drug reactions and burns. The polypeptide is useful for diagnostic or therapeutic purposes. The binding molecules are also useful for pretargeting applications for chemotherapeutic drug delivery. Dwg.0/35

ABEX

UPTX: 20050126

ADMINISTRATION - Dosage is 1-10 mg/kg body weight administered via parenteral, topical, intravenous, oral, subcutaneous, intraarterial, intracranial, intraperitoneal, intranasal or intramuscular means.

EXAMPLE - No relevant example given.

L15 ANSWER 10 OF 12 WPIX COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER:

2005-058132 [06] WPIX

CROSS REFERENCE: DOC. NO. CPI: 2005-058133 [06]

TITLE:

C2005-020195
New composition comprising polypeptide dimers having at

least two binding sites and at least two polypeptide chains comprising a heavy chain portion and a synthetic peptide, useful for treating e.g., cancer or

autoimmune diseases.

DERWENT CLASS:

B04 D16

LV MC MK NL PL PT RO SE SI SK TR

INVENTOR(S):

BRASLAWSKY, G; CHINN, P; GLASER, S;

HOPP, J; YANG, T; BRASLAWSKY, G R

PATENT ASSIGNEE(S):

(BIOJ) BIOGEN IDEC MA INC

COUNTRY COUNT:

109

PATENT INFORMATION:

PAT	rent	NO			KI	ND I	TAC	Ξ	Ī	WEEI	X		LA	1	PG								
WO	200	5000	0898	3 3	A2	200	0501	L06	(20	0050	06)	* El	<b>J</b> :	152	-								
	RW:	ΑT	BE	BG	${\tt BW}$	CH	CY	CZ	DE	DK	EΑ	EE	ES	FI	FR	GB	GH	GM	GR	HU	ΙE	IT	KE
		LS	LU	MC	MW	MZ	NA	NL	OA	PL	PT	RO	SD	SE	SI	SK	$\mathtt{SL}$	sz	TR	TZ	UG	ZM	zw
	W:	ΑE	AG	AL	AM	AT	ΑU	ΑZ	BA	BB	BG	BR	BW	BY	BZ	CA	CH	CN	CO	CR	CU	CZ	DE
		DK	DM	DZ	EC	EE	EG	ES	FI	GB	GD	GΕ	GH	GM	HR	HU	ID	$_{ m IL}$	IN	IS	JΡ	ΚE	KG
		ΚP	KR	ΚZ	LC	LK	LR	LS	LT	LU	LV	MA	MD	MG	MK	MN	MW	MX	ΜZ	NA	NI	NO	NZ
		OM	PG	PH	PL	PT	RO	RU	SC	SD	SE	SG	SK	$\mathtt{SL}$	SY	ТJ	TM	TN	TR	TT	TZ	UΑ	UG
		US	UZ	VC	VN	YU	ZA	zM	ZW														
US	200	5163	3783	3	<b>A</b> 1	200	0507	728	(20	005	50)												
EP	164	1826	5		A2	200	0604	105	(20	0062	24)	El	N.										
	R:	AL	AΤ	ΒE	BG	CH	CY	CZ	DE	DK	ΕE	ES	FI	FR	GB	GR	HR	HU	ΙE	IT	LI	LT	LU

#### APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 2005000898 US 2005163783	A2 A1 Provisional Provisional Provisional Provisional Provisional	WO 2004-US20944 US 2003-483877P US 2003-508810P US 2003-515351P US 2003-516030P US 2004-880320	20040628 20030627 20031003 20031028 20031030 20040628
EP 1641826	A2	EP 2004-777278 WO 2004-US20944	20040628 20040628

#### FILING DETAILS:

PATENT NO	KIND	PATENT NO
EP 1641826	A2 Based on	WO 2005000898
PRIORITY APPLN.	INFO: US 2003-516030P 2003-483877P 2003-508810P 2003-515351P 2004-880320	20031030; US 20030627; US 20031003; US 20031028; US 20040628

AB WO2005000898 A UPAB: 20060410

NOVELTY - A composition comprising polypeptide dimers having at least two binding sites and at least two polypeptide chains, where the polypeptide chains comprise at least one heavy chain portion and a synthetic connecting peptide, is new.

DETAILED DESCRIPTION - A composition comprising polypeptide dimers having at least two binding sites and at least two polypeptide chains, where the polypeptide chains comprise at least one heavy chain portion and a synthetic connecting **peptide** where greater than 50% of the polypeptide dimers comprise polypeptide chains that are linked via at least one interchain disulfide linkage, and where the connecting **peptide** comprises a proline residue at position 243 of the Kabat numbering system, is new.

INDEPENDENT CLAIMS are also included for:

- (1) a nucleic acid molecule comprising a nucleotide sequence encoding a polypeptide chain above, where the nucleic acid molecule comprises a nucleotide sequence comprising 1044, 1083, 1089, 1095, 1095, 1107 or 368 bp (SEQ ID NOs:16, 20,21, 38, 42, 46 or 47, respectively) fully defined in the specification, or comprising a nucleotide sequence comprising 1089 or 663 bp (SEQ ID NOs: 24 or 25, respectively) fully defined in the specification;
  - (2) a host cell comprising the nucleic acid molecule of (1);
- (3) a connecting peptide comprising or consisting of the amino acid sequence selected from: Glu-Pro-Lys-Ser-Cys-Asp-Lys-Thr-His-Thr-Cys-Pro-Pro-Cys-Pro-Glu-Pro-Lys-Ser-Cys-Asp-Thr-Pro-Pro-Pro-Cys-Pro-Arg-Cys-Pro-Gly-Gly-Gly-Ser-Ser-Gly-Gly-Gly-Ser-Gly (SEQ ID NO: 8), Glu-Pro-Lys-Ser-Cys-Asp-Lys-Thr-His-Thr-Cys-Pro-Pro-Cys-Pro-Glu-Pro-Lys-Ser-Cys-Asp-Thr-Pro-Pro-Pro-Cys-Pro-Arg-Cys-Pro-Arg-Pro-Gly-Gly-Gly-Ser-Ser-Gly-Gly-Gly-Ser-Gly (SEQ ID NO: 9), Glu-Pro-Lys-Ser-Cys-Asp-Lys-Thr-His-Thr-Ser-Pro-Cys-Pro-Gly-Gly-Gly-Ser-Ser-Gly-Gly-Ser-Gly (SEQ ID NO: 10), Glu-Pro-Lys-Ser-Cys-Asp-Lys-Thr-His-Thr-Ser-Pro-Pro-Cys-Pro-Ala-Pro-Gly-Gly-Gly-Ser-Ser-Gly-Gly-Ser-Gly (SEQ ID NO: 11), Glu-Pro-Lys-Ser-Cys-Asp-Lys-Thr-His-Thr-Cys-Pro-Pro-Ser-Pro-Gly-Gly-Gly-Ser-Ser-Gly-Gly-Gly-Ser-Gly (SEQ ID NO: 12), Glu-Pro-Lys-Ser-Cys-Asp-Lys-Thr-His-Thr-Cys-Pro-Pro-Ser-Pro-Ala-Pro-Gly-Gly-Gly-Ser-Ser-Gly-Gly-Ser-Gly (SEQ ID NO: 13), Glu-Pro-Lys-Ser-Cys-Asp-Lys-Thr-His-Thr-Cys-Pro-Pro-Cys-Pro-Ala-Pro-Gly-Gly-Gly-Ser-Ser-Gly-Gly-Gly-Ser-Gly (SEQ ID NO: 14), Glu-Pro-Lys-Ser-Cys-Asp-Lys-Thr-His-Thr-Cys-Pro-Pro-Cys-Pro-Gly-Gly-Gly-Ser-Ser-Gly-Gly-Gly-Ser-Gly (SEQ ID NO: 15) and Glu-Ser-Lys-Tyr-Gly-Pro-Pro-Cys-Pro-Ser-Cys-Pro-Glu-Pro-Lys-Ser-Cys-Asp-Thr-Pro-Pro-Pro-Cys-Pro-Arg-Cys-Pro-Ala-Pro (SEQ ID NO: 53);
- (4) a domain deleted antibody molecule comprising an amino acid sequence comprising 347, 360, 362, 362, 220, 365 or 365 bp (SEQ ID NOs: 18, 22, 23, 26, 27, 40 or 44, respectively) fully defined in the specification;
- (5) an antibody molecule comprising the amino acid sequence comprising 113 or 115 amino acids (SEQ ID NO: 31 or 35, respectively) fully defined in the specification;
  - (6) separating a first and a second polypeptide dimer where the first

polypeptide dimer comprises at least two binding sites and at least two polypeptide chains, where the polypeptide chains comprises a heavy chain portion and where the first polypeptide dimer comprises polypeptide chains that are linked via at least one disulfide linkage and where the second polypeptide dimer comprises at least two binding sites and at least two polypeptide chains, where the polypeptide chains comprises a heavy chain portion and where the second polypeptide dimer comprises polypeptide chains that are not linked via at least one disulfide linkage;

- (7) separating a first properly folded antibody molecule from a second improperly folded antibody molecule, where each of the first and second antibody molecules comprises four polypeptide chains, where at least two of the chains comprise at least one heavy chain portion, and at least two of the chains comprise at least one light chain portion;
- (8) increasing the amount of a first polypeptide dimer relative to the amount of a second polypeptide dimer produced by a cell, where the first and second polypeptide dimers comprise at least two binding sites and at least two polypeptide chains, the polypeptide chains comprising a heavy chain portion, where the first dimer comprises polypeptide chains that are linked via at least one disulfide linkage and where the second dimer comprises polypeptide chains that are not linked via at least one disulfide linkage;
- (9) a composition comprising a first polypeptide dimer prepared by the method of (4), or comprising a first polypeptide prepared by the method of (5), or made by the method of (6);
- (10) a polypeptide comprising a synthetic connecting **peptide** which comprises the amino acid sequence Cys-Pro-Glu-Pro-Lys-Ser-Cys-Asp-Thr-Pro-Pro-Cys-Pro-Arg (SEQ ID NO: 37), where the polypeptide is not a naturally occurring IgG3 molecule; and
- (11) increasing the amount of dimers comprising polypeptide chains linked via at least one disulfide linkage in a population of IgG4 molecules produced by a cell.

ACTIVITY - Immunosuppressive; Antianemic; Dermatological; Muscular-Gen.; Neuroprotective; Thyromimetic; Antithyroid; Nephrotropic; Antirheumatic; Antiarthritic; Antiinflammatory; Antiulcer; Gastrointestinal-Gen. No biological data given.

MECHANISM OF ACTION - None given.

USE - The composition is useful for treating a subject that would benefit from treatment with a binding molecule, where the subject is suffering from cancer, lymphoma, an autoimmune disease or disorder, or inflammatory disease or disorder (claimed). The composition is useful for treating autoimmune diseases such as Crohn's disease, inflammatory bowel disease, systemic lupus erythematosus, ulcerative colitis, rheumatoid arthritis, Goodpasture's syndrome, Grave's disease, Hashimoto's thyroiditis, pemphigus vulgaris, myasthenia gravis, scleroderma, autoimmune hemolytic anemia, pernicious anemia, and Sjogren's syndrome. Dwg.0/35

ABEX

UPTX: 20050126

ADMINISTRATION - Dosage is 1-10 mg/kg body weight administered via parenteral, topical, intravenous, oral, subcutaneous, intraarterial, intracranial, intraperitoneal, intranasal or intramuscular means.

EXAMPLE - No relevant example given.

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L15 ANSWER 11 OF 12 WPIX COPYRIGHT 2006 THE THOMSON CORP on STN ACCESSION NUMBER: 2002-682677 [73] WPIX
CROSS REFERENCE: 1997-108638 [10]; 1998-286601 [25]; 2001-335883 [35]; 2002-089895 [12]; 2002-154869 [20]; 2002-619209 [66]; 2003-441463 [41]
DOC. NO. CPI: C2002-192540
TITLE: Use of CD23 antagonists for inducing apoptosis in
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malignant cells, or treating neoplastic disorders in a mammal, e.g. resistant Hodgkin's disease, Burkitt's lymphoma or B cell chronic lymphocytic leukemia.

DERWENT CLASS:

B02 B04 D16 K08

W 20050210 (200511)

A 20050223 (200519)#

INVENTOR (S):

BRASLAWSKY, G R; HANNA, N; HARIHARAN, K;

PATHAN, N; BRASLAWSKY, G

PATENT ASSIGNEE(S):

(IDEC-N) IDEC PHARM CORP; (BRAS-I) BRASLAWSKY G; (HANN-I)

138

94

HANNA N; (HARI-I) HARIHARAN K; (PATH-I) PATHAN N

COUNTRY COUNT: 10

PATENT INFORMATION:

PAT	TENT	ИО			KI	MD I	DATE	Ξ	V	VEE	ζ.		LA	I	PG								
WO	2002	2060	0484	· 1	A1	200	0208	308	(20	002	73);	El	J	88	•								
	RW:	ΑT	BE	CH	CY	DE	DK	EΑ	ES	FI	FR	GB	GH	GM	GR	ΙE	IT	KE	LS	LU	MC	MW	MZ
		NL	OA	PT	SD	SE	$\mathtt{SL}$	SZ	TR	TZ	UG	ZM	zw										
	W:	ΑE	AG	AL	ΑM	ΑT	ΑU	AZ	ва	вв	ВG	BR	BY	BZ	CA	CH	CN	CO	CR	CU	CZ	DE	DK
		DM	DZ	EC	EE	ES	FΙ	GB	GD	GE	GH	GM	HR	HU	ID	IL	IN	IS	JΡ	KE	KG	ΚP	KR
		ΚZ	LC	LK	LR	LS	LT	LU	LV	MA	MD	MG	MK	MN	MW	MX	ΜZ	NO	NZ	MO	PH	PL	PT
		RO	RU	SD	SE	SG	SI	SK	$\mathtt{SL}$	ТJ	TM	TN	TR	TT	TZ	UA	UG	US	UZ	VN	YU	ZA	ZM
		ZW																					
US	200	215	9996	5	<b>A</b> 1	200	210	31	(20	002	74)												
NO	2003	300	3417	7	Α	200	0309	930	(20	003	73)												
EP	1370	029	2		<b>A</b> 1	200	312	217	(20	040	02)	Eì	V										
	R:	AL	ΑT	BE	CH	CY	DE	DK	ES	FI	FR	GB	GR	ΙE	IT	LI	LT	LU	LV	MC	MK	NL	PT
		RO	SE	SI	TR																		
AU	2002	223	7972	2	<b>A</b> 1	200	208	312	(20	004	27)												

#### APPLICATION DETAILS:

JP 2005503999

ZA 2003005891

PATENT NO	KIND	APPLICATION	DATE
WO 2002060484	A1	WO 2002-US2620	20020131
US 2002159996	A1 CIP of	US 2001-772938	20010131
		US 2001-985646	20011105
NO 2003003417	Α	WO 2002-US2620	20020131
		NO 2003-3417	20030730
EP 1370292	A1	EP 2002-704280	20020131
		WO 2002-US2620	20020131
AU 2002237972	A1	AU 2002-237972	20020131
JP 2005503999	W	JP 2002-560675	20020131
		WO 2002-US2620	20020131
ZA 2003005891	A	ZA 2003-5891	20030730

#### FILING DETAILS:

PATENT NO	KIND	PATENT NO
EP 1370292 AU 2002237972 JP 2005503999	Al Based on Al Based on W Based on	WO 2002060484 WO 2002060484 WO 2002060484
PRIORITY APPLN. INFO	: US 2001-985646 2001-772938 2001-855717 2003-5891	20011105; US 20010131; US 20010516; ZA 20030730

AB WO 200260484 A UPAB: 20051130

NOVELTY - Treating a neoplastic disorder in a mammal comprises

10/796,158

administering a CD23 antagonist to the mammal.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) inducing apoptosis (M1) in malignant cells comprising contacting the malignant cells with a CD23 antagonist; and
- (2) kit useful for treating a mammal suffering from or predisposed to a neoplastic disorder comprising:
- (a) at least one container having a CD23 antagonist deposited in it;
- (b) label or insert indicating that the CD23 antagonist may be used to treat the neoplastic disorder.

ACTIVITY - Cytostatic.

 ${\tt MECHANISM}$  OF ACTION - CD23-Antagonist; Synergist; Inducer of Apoptosis.

Anti-CD23 antibodies were examined to determined to what extent they can induce apoptosis in malignant cells. Apoptosis was measured by a caspase-3 activation assay. Percent apoptosis was documented at 4 and 24 hours using mean fluorescent intensity in log scale (MFI). SKW cells grown in the presence of IDEC-152 (p5E8) did not show substantial activation of caspase-3 (3.8% apoptosis after 4 hours, 3.65% after 24 hours). However, cross-linking IDEC-152 and Rituxan on the SKW cell surface resulted in increased activation of caspase-3 (80.26% and 78.5% apoptosis after 4 hours, 60.51% and 66.49% after 24 hours). By comparison, cultures added with the isotype matched control antibody (CE9.1) of irrelevant specificity did not show any apoptosis. Thus, the results showed that IDEC-152 mediated antibody-dependent cell-mediated cytotoxicity (ADCC) activity of tumor cells, and induced apoptosis in CD23+ tumor cells.

USE - The method is useful for treating a neoplastic disorder in a mammal, or inducing apoptosis in malignant cells (claimed). The neoplastic disorder includes relapsed Hodgkin's disease; resistant Hodgkin's disease; high grade, low grade and intermediate grade non-Hodgkin's lymphomas; lymphoplasmacytoid lymphoma (LPL); mantle cell lymphoma (MCL); follicular lymphoma (FL); diffuse large cell lymphoma (DLCL); Burkitt's lymphoma (BL); AIDS-related lymphomas; monocytic B cell lymphoma; angioimmunoblastic lymphoadenopathy; small lymphocytic; follicular, diffuse large cell; diffuse small cleaved cell; large cell immunoblastic lymphoblastoma; small, non-cleaved; Burkitt's or non-Burkitt's; follicular, predominantly large cell; follicular, predominantly small cleaved cell; follicular, mixed small and large cell lymphomas; or particularly B cell chronic lymphocytic leukemia (B-CLL) (claimed).

ABEX UPTX: 20021113

ADMINISTRATION - Dosage is 0.05-100, preferably 0.5-10, mg/kg body weight per day. Administration is oral, parenteral (e.g. intravenous, intraarterial, intraperitoneal, intramuscular, subcutaneous, rectal or vaginal), by inhalation or topical. The CD23 antagonist and chemotherapeutic agent (preferably an antibody) may be administered in any order or concurrently.

L15 ANSWER 12 OF 12 WPIX COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 1999-357351 [30] WPIX

DOC. NO. CPI: C1999-105653

TITLE: New immunogenic compositions for treating cancer or virus

or parasite infection.

DERWENT CLASS: A96 B04 D16

INVENTOR(S): BRASLAWSKY, G R; HANNA, N; HARIHARAN, K;

HARIHARA, K

PATENT ASSIGNEE(S): (IDEC-N) IDEC PHARM CORP; (BIOG-N) BIOGEN IDEC INC;

(BIOJ) BIOGEN IDEC INC

COUNTRY COUNT: 84

#### PATENT INFORMATION:

PA	rent	NO			KI	ND I	TAC	3	V	WEE!	К		LA	I	PG								
WO	991:	3912	- <b></b> -		A1	19:	990:	325	(19	999:	30)	* Eì	J	41									
	RW:	AT	BE	СН	CY	DE	DK	EA	ES	FI	FR	GB	GH	GM	GR	ΙE	IT	ΚE	LS	LU	MC	MW	NL
		OA	PT	SD	SE	sz	UG	ZW															
	W:	AL	AΜ	ΑT	ΑU	AZ	BA	BB	ВG	BR	BY	CA	CH	CN	CU	CZ	DE	DK	EE	ES	FΙ	GB	GE
		GH	GM	HR	HU	ID	IL	IS	JР	KE	KG	ΚP	KR	ΚZ	LC	LK	LR	LS	LT	LU	$r_{\Lambda}$	MD	MG
						ИО	ΝZ	PL	PT	RO	RU	SD	SE	SG	SI	SK	$s_L$	ТJ	TM	TR	TT	UA	UG
			VN																				
	980													36									
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EP	101!	-																					
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	221	_							•														
	2004								•		•												
US	6998	312	5		В2	200	0602	214	(20	006	13)												

#### APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 9913912	A1	WO 1998-US18495	19980917
ZA 9808461	Α	ZA 1998-8461	19980916
AU 9895658	A	AU 1998-95658	19980917
EP 1015031	A1	EP 1998-949313	19980917
		WO 1998-US18495	19980917
NO 2000001413	A	WO 1998-US18495	19980917
		NO 2000-1413	20000317
CN 1279616	A	CN 1998-811280	19980917
US 2001018054	Al Cont of	US 1997-933359	19970918
		US 2001-853580	20010514
US 2001019715	Al Div ex	US 1997-933359	19970918
		US 2001-853581	20010514
KR 2001024109	A	KR 2000-702864	20000317
JP 2001516727	W	WO 1998-US18495	19980917
		JP 2000-511527	19980917
AU 742216	В	AU 1998-95658	19980917
RU 2219947	C2	WO 1998-US18495	19980917
		RU 2000-109595	19980917
US 2004137014	A1 Div ex	US 1997-933359	19970918
	Div ex	US 2001-853581	20010514
		US 2003-743739	20031224
US 6998125	B2 Div ex	US 1997-933359	19970918
		US 2001-853581	20010514

#### FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 9895658	A Based on	WO 9913912
EP 1015031	Al Based on	WO 9913912

JP 2001516727 W Based on WO 9913912
AU 742216 B Previous Publ. AU 9895658
Based on WO 9913912
RU 2219947 C2 Based on WO 9913912

PRIORITY APPLN. INFO: US 1997-933359 19970918; US 2001-853580 20010514; US 2001-853581 20010514; US

2003-743739 20031224

AB WO 9913912 A UPAB: 19990802

NOVELTY - New immunogenic compositions for treating cancer or virus or parasite infection comprise a combination of antigen formulation and an agent capable of neutralizing or down-regulating immunosuppressive factors.

DETAILED DESCRIPTION - A composition (A) comprises:

- (a) an admixture comprising a cancer, viral or parasitic antigen expressed by cancer, virally or parasitic infected cells and a microfluidized antigen formulation (MAF) (formulated as a stable oil-in-water emulsion), the antigen formulation comprising:
  - (i) a stabilizing detergent;
  - (ii) a micelle-forming agent; and
  - (iii) a biodegradable and biocompatible oil; and
- (b) at least one agent which is capable of neutralizing or down-regulating the activity of immunosuppressive factors.

INDEPENDENT CLAIMS are also included for the following:

- (1) a method of treatment which includes the induction of a cytotoxic T-lymphocyte (CTL) response where the improvement comprises:
- (a) the administration of an adjuvant which induces a CTL response;
- (b) the administration of an antagonist of an immunosuppressive factor, where the administration of adjuvant and antagonist is effected sequentially or concurrently, and in any order;
- (2) a method of restoring or boosting hematopoiesis comprising administering to a patient:
- (a) an admixture as in (A) (a) which is administered to the patient to induce a CTL response in the patient which is specific for the viral or cancer antigen contained in the admixture; and
- (b) at least one agent which is capable of neutralizing or down regulating the activity of tumor and host secreted immunosuppressive factors, where the admixture and the agent are administered separately or in combination, and in any order;
- (3) a composition comprising an admixture as in (A) (a) and one or more transforming growth factor (TGF) beta antagonists;
  - (4) treatment of neoplastic or cancerous growths, comprising:
- (a) administration of an admixture comprising a cancer or tumor antigen expressed by the cancer cells and a MAF (described above); and
- (b) administration of at least one agent which is capable of neutralizing or down-regulating the activity of tumors and host secreted immunosuppressive factors. The admixture is administered in an amount sufficient to induce a cytotoxic T-lymphocyte response in the patient which is specific for the cancer or tumor antigen contained in the admixture.

ACTIVITY - Antitumor; Antiviral; Antiparasitic.

MECHANISM OF ACTION - Induction of a cytotoxic T-lymphocyte response. USE - The methods can be used for restoring or boosting hematopoiesis (claimed). They can be used for treating cancers, e.g. breast cancer, brain cancer, cervical cancer, leukemia, lymphoma, prostate cancer, skin cancer, bladder cancer, kidney cancer, myeloma, colorectal cancer, or endometrial cancer, viral infections e.g. papillomavirus, hepatitis, herpes, cytomegalovirus, respiratory syncytial virus or HIV, or parasitic

infection, e.g. malaria (claimed). The agent which is capable of neutralizing or down-regulating the activity of immunosuppressive factors enhances the efficacy of tumor/viral vaccines.

ADVANTAGE - The combinations of the antigen compositions and antagonists of immunosuppressive agents results in a synergistic enhancement of CTL response, thereby resulting in enhanced therapeutic response against targeted antigen-expressing cells.

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ABEX

UPTX: 19990802

ADMINSTRATION - Administration of the adjuvant is especially intradermal, intramuscular or subcutaneous. Administration of the TGF antagonist is especially intravenous.

EXAMPLE - Mice were inoculated with ovalbumin expressing EG7 cells (2 x 10 to the power 6 cells/mouse). On day 7, post-inoculation mice bearing 250-350 mm3 size tumors were sorted into groups. Group A, the control group received no antigen injection. Group B received 30 microg of ovalbumin in PROVAX (RTM) s.c. Group C received 30 microg of ovalbumin in PROVAX (RTM) s.c. and 50 microg of anti-transforming growth factor beta (TGF-beta) antibodies i.p. Per mouse. Group D received 50 microg of anti-TGF beta antibodies i.p.

The results showed that the treatment of mice bearing progressively growing EG7 tumors with anti-TGF-beta antibodies in conjunction with ovalbumin in PROVAX gave enhanced anti-tumor activity under conditions where treatment with ovalbumin-PROVAX (RTM) is not effective.

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L14

(FILE 'HOME' ENTERED AT 16:01:15 ON 06 JUN 2006)

FILE 'REGISTRY' ENTERED AT 16:01:20 ON 06 JUN 2006 6 SEA ABB=ON PLU=ON YCYYCFWKTCT | CYYYCFWKTCT | YYCYCFWKTCT/SQSP L1

FILE 'CAPLUS' ENTERED AT 16:02:47 ON 06 JUN 2006

1 SEA ABB=ON PLU=ON L1 L2

D AU

E BRASLAWSKY G/AU

E CHINN P/AU

FILE 'REGISTRY' ENTERED AT 16:04:18 ON 06 JUN 2006

D QUE L1

D L1 RN CN SQL KWIC NTE

D L1 RN CN SQL KWIC NTE 2-6

FILE 'CAPLUS' ENTERED AT 16:07:50 ON 06 JUN 2006

D QUE L2

D IBIB ED AB HITRN

D SCAN L2

41 SEA ABB=ON PLU=ON BRASLAWSKY G?/AU L3

21 SEA ABB=ON PLU=ON CHINN P?/AU L4

5 SEA ABB=ON PLU=ON (L3 OR L4) AND PEPTIDES/CW L5

FILE 'MEDLINE' ENTERED AT 16:20:49 ON 06 JUN 2006

L6 35 SEA ABB=ON PLU=ON BRASLAWSKY G?/AU

92 SEA ABB=ON PLU=ON CHINN P?/AU L7

3 SEA ABB=ON PLU=ON (L3 OR L4) AND PEPTIDE? L8

D TRIAL 1-3

FILE 'EMBASE' ENTERED AT 16:21:24 ON 06 JUN 2006

25 SEA ABB=ON PLU=ON BRASLAWSKY G?/AU L9

18 SEA ABB=ON PLU=ON CHINN P?/AU L10

1 SEA ABB=ON PLU=ON L3 AND L4 L11

2 SEA ABB=ON PLU=ON (L3 OR L4) AND PEPTIDE? L12

FILE 'BIOSIS' ENTERED AT 16:25:01 ON 06 JUN 2006

L13 2 SEA ABB=ON PLU=ON (L3 OR L4) AND PEPTIDE?

FILE 'WPIX' ENTERED AT 16:25:15 ON 06 JUN 2006

7 SEA ABB=ON PLU=ON (L3 OR L4) AND PEPTIDE?

FILE 'CAPLUS' ENTERED AT 16:26:43 ON 06 JUN 2006 D QUE L5

FILE 'MEDLINE' ENTERED AT 16:27:01 ON 06 JUN 2006 D QUE L8

FILE 'EMBASE' ENTERED AT 16:27:08 ON 06 JUN 2006 D QUE L12

FILE 'BIOSIS' ENTERED AT 16:27:15 ON 06 JUN 2006 D QUE L13

FILE 'WPIX' ENTERED AT 16:27:21 ON 06 JUN 2006 D OUE L14

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D IBIB AB ABEX L15 9-12

FILE 'MEDLINE, CAPLUS, EMBASE, BIOSIS, WPIX' ENTERED AT 16:27:36 ON 06 JUN 2006

L15

12 DUP REM L8 L5 L12 L13 L14 (7 DUPLICATES REMOVED)

ANSWERS '1-3' FROM FILE MEDLINE

ANSWERS '4-8' FROM FILE CAPLUS

ANSWERS '9-12' FROM FILE WPIX

D IBIB ED AB L15 1-8

FILE HOME

FILE REGISTRY

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 5 JUN 2006 HIGHEST RN 886840-90-0 DICTIONARY FILE UPDATES: 5 JUN 2006 HIGHEST RN 886840-90-0

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH January 6, 2006

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Structure search iteration limits have been increased. See HELP SLIMITS for details.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

http://www.cas.org/ONLINE/UG/regprops.html

FILE CAPLUS

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FILE COVERS 1907 - 6 Jun 2006 VOL 144 ISS 24 FILE LAST UPDATED: 5 Jun 2006 (20060605/ED) Effective October 17, 2005, revised CAS Information Use Policies apply. They are available for your review at:

http://www.cas.org/infopolicy.html

#### FILE MEDLINE

FILE LAST UPDATED: 3 JUN 2006 (20060603/UP). FILE COVERS 1950 TO DATE.

On December 11, 2005, the 2006 MeSH terms were loaded.

The MEDLINE reload for 2006 is now (26 Feb.) available. For details on the 2006 reload, enter HELP RLOAD at an arrow prompt (=>). See also:

http://www.nlm.nih.gov/mesh/

http://www.nlm.nih.gov/pubs/techbull/nd04/nd04 mesh.html

http://www.nlm.nih.gov/pubs/techbull/nd05/nd05\_med\_data\_changes.html

http://www.nlm.nih.gov/pubs/techbull/nd05/nd05 2006 MeSH.html

OLDMEDLINE is covered back to 1950.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2006 vocabulary.

This file contains CAS Registry Numbers for easy and accurate substance identification.

#### FILE EMBASE

FILE COVERS 1974 TO 6 Jun 2006 (20060606/ED)

EMBASE has been reloaded. Enter HELP RLOAD for details.

EMBASE is now updated daily. SDI frequency remains weekly (default) and biweekly.

This file contains CAS Registry Numbers for easy and accurate substance identification.

#### FILE BIOSIS

FILE COVERS 1969 TO DATE.

CAS REGISTRY NUMBERS AND CHEMICAL NAMES (CNs) PRESENT FROM JANUARY 1969 TO DATE.

RECORDS LAST ADDED: 31 May 2006 (20060531/ED)

#### FILE WPIX

FILE LAST UPDATED: 2 JUN 2006 <20060602/UP>
MOST RECENT DERWENT UPDATE: 200635 <200635/DW>
DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE

>>> FOR A COPY OF THE DERWENT WORLD PATENTS INDEX STN USER GUIDE, PLEASE VISIT:

http://www.stn-international.de/training\_center/patents/stn\_guide.pdf <

>>> FOR DETAILS OF THE PATENTS COVERED IN CURRENT UPDATES, SEE http://scientific.thomson.com/support/patents/coverage/latestupdates/

>>> PLEASE BE AWARE OF THE NEW IPC REFORM IN 2006, SEE http://www.stn-international.de/stndatabases/details/ipc\_reform.html and

http://scientific.thomson.com/media/scpdf/ipcrdwpi.pdf <<<

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